Patent Application Attorney Docket No. 10925A

## **REMARKS**

Claims 1 – 37 were submitted with the original specification of this application. Applicants have hereinabove canceled claims 10-18 and 23-37 without prejudice to applicants' right to pursue the subject matter of the canceled claims in one or more divisional or continuation applications, and have amended claims 1-9 and 19-22 and added new claims 38-56. Accordingly, upon entry of this amendment, claims 1-9, 19-22 and 38-56 are pending.

Applicants maintain that the amendment of claims 1-9 and 19-22, and the addition of new claims 38-56 raises no issue of new matter. All of the amendments and additions are made to conform the claims to U.S. practice and to reduce the number of independent claims so as to reduce excess claims fees. Claims 23 to 37 have been replaced with new claims 38-56, with the correspondence indicated below:

- New claim 38 corresponds to original claim 24;
- New claims 39, 41, 43, 50, 52 and 54 correspond to original claim 31;
- New claim 40 corresponds to original claim 23;
- New claim 42 corresponds to original claim 25;
- New claim 44 corresponds to original claim 33;
- New claim 45 corresponds to original claim 27;
- New claim 46 corresponds to original claim 32;
- New claim 47 corresponds to original claim 26;
- New claim 48 is new, and was added to properly claim a preferred embodiment of the invention of original claim 26;
- New claim 49 corresponds to original claim 28;
- New claim 51 corresponds to original claim 29;
- New claim 53 corresponds to original claim 30;
- New claim 55 corresponds to original claims 34, 35 and 36; and
- New claim 56 corresponds to original claim 37.

Applicants submit that the differences between the new claims and the original claims merely reflect rewording of the claims to U.S. practice, to eliminate multiple dependencies, and to reduce the filing fee by rewriting some independent claims in dependent form. Support for new claim 24 may be found in original claim 26. Accordingly, applicants respectfully request entry of the replacement claims.

Attached hereto is a marked-up version of original claims 1-9 and 9-22 showing the changes made, captioned "<u>PLEASE DO NOT ENTER - Version with markings to show changes made</u>."

The filing fee has been calculated on the basis of the replacement claims. No additional fee is believed necessary in connection with this amendment. However, the fee transmittal submitted herewith authorizes payment of any additional fees.



Applicants earnestly solicit favorable action on the merits.

Respectfully submitted,

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Patent Application

Guide to claim amendments:

Insertions – underlined

Deletions – strikethrough

1. (Amended) A compound of the formula:

$$O = \begin{pmatrix} R^1 & H_3C & CH_3 \\ NH & N & N \\ N & N & N \\ N & N & N \\ H_3C & N & N \\ \end{pmatrix}$$

or a pharmaceutically acceptable salt or solvate thereof, wherein:

 $R^1$  is  $C_{3-6}$  cycloalkyl optionally substituted by one or more fluorine atoms, or  $C_{1-6}$  alkyl optionally substituted by one or more fluorine atoms, or  $C_{3-6}$  cycloalkylmethyl optionally ring-substituted by one or more fluorine atoms;

and

R<sup>2</sup> is phenyl optionally substituted by one or more fluorine atoms: or a pharmaceutically acceptable salt or solvate thereof.

2. (Amended) AThe compound as claimed inof claim 1 of the formula:

$$O \longrightarrow NH \longrightarrow H_3C \longrightarrow CH_3$$

$$IA \longrightarrow N \longrightarrow N$$

$$H_3C \longrightarrow N$$

or a pharmaceutically acceptable salt or solvate thereof, wherein:

- $-R^1$  represents is either  $C_{3-6}$  cycloalkyl optionally substituted by one or more fluorine atoms, or  $C_{1-6}$  alkyl optionally substituted by one or more fluorine atoms, or a pharmaceutically acceptable salt or solvate thereof.
- 3. (Amended) A<u>The</u> compound as claimed inof claim 1, wherein R<sup>1</sup> is either  $C_{4-6}$  cycloalkyl optionally substituted by one or two fluorine atoms, or  $C_{1-4}$  alkyl optionally substituted by from one to three fluorine atoms.

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- 4. (Amended) AThe compound as claimed inof claim 3, wherein R<sup>1</sup> is either cyclobutyl, cyclopentyl, 4,4-difluorocyclohexyl or 3,3,3-trifluoropropyl.
- 5. (Amended) A<u>The</u> compound as claimed inof claim 1, 3 or 4 wherein R<sup>2</sup> is phenyl optionally substituted by 1 or 2 fluorine atom(s).
- 6. (Amended) AThe compound as claimed inof claim 5, wherein R<sup>2</sup> is phenyl or monofluorophenyl.
- 7. (Amended) A<u>The</u> compound as claimed inof claim 6, wherein R<sup>2</sup> is phenyl or 3-fluorophenyl.
- 8. (Amended) AThe compound as claimed inof claim 1 which is selected from the group consisting of:

N-{(1S)-3-[3-(3-lsopropyl-5-methyl-4H-1,2,4-triazol-4-yl)-exo-8-azabicyclo[3.2.1]oct-8-yl]-1-phenylpropyl}cyclobutanecarboxamide;

N-{(1S)-3-[3-(3-Isopropyl-5-methyl-4H-1,2,4-triazol-4-yl)-exo-8-

azabicyclo[3.2.1]oct-8-yl]-1-phenylpropyl}cyclopentanecarboxamide;

N-{(1S)-3-[3-(3-lsopropyl-5-methyl-4H-1,2,4-triazol-4-yl)-exo-8-

azabicyclo[3.2.1]oct-8-yl]-1-phenylpropyl}-4,4,4-trifluorobutanamide;

N-{(1S)-3-[3-(3-Isopropyl-5-methyl-4H-1,2,4-triazol-4-yl)-exo-8-

azabicyclo[3.2.1]oct-8-yl]-1-phenylpropyl}-4,4-difluorocyclohexanecarboxamide; and

N-{(1S)-3-[3-(3-lsopropyl-5-methyl-4H-1,2,4-triazol-4-yl)-exo-8-

azabicyclo[3.2.1]oct-8-yl]-1-(3-fluorophenyl)propyl}-4,4-

difluorocyclohexanecarboxamide:;

or a pharmaceutically acceptable salt or solvate of any thereof.

- 9. (Amended) A pharmaceutical composition including comprising a compound of the formula (I) or a pharmaceutically acceptable salt or solvate thereof, as claimed in any preceding claim 1, together with and one of a pharmaceutically acceptable excipient, a pharmaceutically acceptable diluent or a pharmaceutically acceptable carrier.
- 19. (Amended) A method of treatingment of in a mammal to treat a disorder in which the modulation of CCR5 receptors is implicated, including treatingwhich comprises administering to said mammal with an effective amount of a compound of the formula (I)

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or with a pharmaceutically acceptable salt, solvate or composition thereof as claimed in any one of claims 1 to 8 and 9, respectively.

- 20. (Amended) A method of treatingment of a mammal to treat HIV, a retroviral infection genetically related to HIV, AIDS, or an inflammatory disease, in a mammal, which comprises administering to including treating-said mammal with an effective amount of a compound of the formula (I) or with a pharmaceutically acceptable salt, solvate or composition thereof as claimed in any one of claims 1 to 8 and 9, respectively.
- 21. (Amended) A method of treating, in a mammal, ment of a mammal to treat a respiratory disorder including selected from adult respiratory distress syndrome (ARDS), bronchitis, chronic bronchitis, chronic obstructive pulmonary disease, cystic fibrosis, asthma, emphysema, rhinitis erand chronic sinusitis, including treating which comprises administering to said mammal with an effective amount of a compound of the formula (I) or with a pharmaceutically acceptable salt, solvate or composition thereof as claimed in any one of claims 1-to 8 and 9, respectively.
- 22. (Amended) A method of treating, in a mammal, ment of a mammal to treat an inflammatory bowel disease, including Crohn's disease or ulcerative colitis, multiple sclerosis, rheumatoid arthritis, graft rejection, including a kidney or a lung allograft, endometriosis, type I diabetes, a renal disease, chronic pancreatitis, an inflammatory lung condition or chronic heart failure including treating which comprises administering to said mammal with an effective amount of a compound of the formula (I) or with a pharmaceutically acceptable salt, solvate or composition thereof as claimed in any one of claims 1 to 8 and 9, respectively.